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PHASE 3 SUMMARY OF MRID 00094000:

PRIMARY EYE IRRITATION IN THE RABBIT

STUDY # 6818A

FLUMETRALIN

GUIDELINE REFERENCE:

81-4 PRIMARY EYE IRRITATION IN THE RABBIT

SUMMARY PREPARED BY:

JACQUELINE GILLIS, Ph.D.

MERRILL TISDEL

14 SEPTEMBER 1990

ORIGINAL STUDY PREPARED BY:

FOOD AND DRUG RESEARCH LABORATORIES, INC.

WAVERLY, NEW YORK

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STATEMENT OF NO DATA CONFIDENTIALITY CLAIMS

No claim of confidentiality is made for any information contained in this study on the basis of its falling within the scope of FIFRA §10(d)(1)(A), (B), or (C).

Company: CIBA-GEIGY Corporation (Typed Name)

Company Agent: Thomas Parshley (Typed Name)

Title: Senior Reg. Specialist

Signature: _____ Date: _____

These data are the property of the Agricultural Division of CIBA-GEIGY Corporation, and as such, are considered to be confidential for all purposes other than compliance with FIFRA §10. Submission of these data in compliance with FIFRA does not constitute a waiver of any right to confidentiality which may exist under any other statute or in any other country.

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FDRL Study No. 6818A

Primary Eye Irritation Study in Rabbits
of CGA-41065 Technical

GLP Compliance Statement

I hereby certify that this study was performed in compliance with regulations for Good Laboratory Practice (GLP) as described by FDA (21 CFR Part 58) and although completed and reported prior to promulgation of the EPA GLP, essentially in compliance with EPA (40 CFR Part 160).

James Lafeglia, Ph.D., President
for Study Director

9/6/90
Date

This study does not meet the requirements for 40 CFR Part 160 since it was conducted prior to the issuance of the EPA Good Laboratory Practice Standards. It was conducted according to the FDA Good Laboratory Practice Standards as indicated above.

Submitter/Sponsor of Study:

Merrill Tisdel
Agricultural Division
CIBA-GEIGY Corporation
Greensboro, North Carolina

PM3006899643

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Certification of Availability of Raw Data

I hereby certify that the submitter possesses or has access to the raw data used in or generated by the study summarized in this document.

Submitter's Representative:

Signature/Date: Merrill Tisdel 10.15.90

Typed Name: Merrill Tisdel

Title: Toxicologist

Certification of Accuracy of Summary and Adequacy of the Study

I certify, in compliance with FIFRA section 4(e)(1)(A), that this summary accurately represents the data presented in the report(s) of this study cited by MRID, and that this study fully satisfies all pertinent requirements of the OPP Guideline it addresses.

Submitter's Representative:

Signature/Date: Merrill Tisdel 10.15.90

Typed Name: Merrill Tisdel

Title: Toxicologist

R406MT0628MG

81-4 Primary Eye Irritation in the Rabbit

ACCEPTANCE CRITERIA

Does your study meet the following acceptance criteria?

1. Y — Technical form of the active ingredient tested. (for reregistration only)
2. N — Study not required if material is corrosive, causes severe dermal irritation or has a pH of ≤ 2 or ≥ 11.5.
3.* Y — 6 adult rabbits
4. Y — Dosing, instillation into the conjunctival sac of one eye per animal.
5.* Y — Dose, 0.1 ml if a liquid; 0.1 ml or not more than 100 mg if a solid, paste or particulate substance.
6. Y — Solid or granular test material ground to a fine dust.
7. Y/N — Eyes not washed for at least 24 hours.
8. Y/N — Eyes examined and graded for irritation before dosing and at 1, 24, 48 and 72 hr, then daily until eyes are normal or 21 days (whichever is shorter).
9.* N — Individual observations for the entire day of dosing.
10.* N — Individual daily observations.

Criteria marked with a * are supplemental and may not be required for every study.

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IDENTIFICATION OF TEST MATERIAL

Chemical Name

CAS Name: N-(2-Chloro-6-fluorobenzyl)-N-ethyl-α,α,α,-trifluoro-2,6-dinitro-p-toluidine

or

2-Chloro-N-[2,6-dinitro-4-(trifluoromethyl)phenyl]-N-ethyl-6-fluorobenzenemethanamine

Common Name: Flumetralin

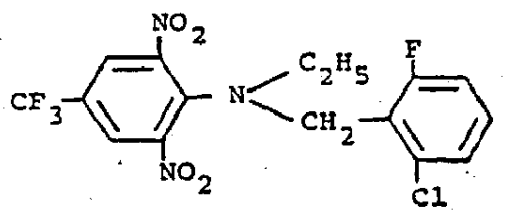
Trade Name: Prime +®

CIBA-GEIGY Code Number: CGA-41065

CAS Registry Number: 62924-70-3

EPA Shaughnessy Number: Unknown

Chemical Structure:



Percent Active Ingredient

92% minimum

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Flumetralin: 81-4: Primary Eye Irritation in the Rabbit

1. The test article was Flumetralin (CGA-41065) Technical, a bright orange crystalline substance, FL-810009, purity 96.4%.

2. The test article is not corrosive, does not cause severe dermal irritation, and does not have a pH less than 2.0 or greater than 11.5.

3. There were six adult male New Zealand white rabbits in the primary 'nonwash' group and three male rabbits in the secondary 'wash' group.

4. The test material was placed into the conjunctival sac of one eye of each animal by gently pulling the lower lid away from the eyeball to form a cup into which the test material was deposited. The lids were then held together for one second.

5. The dose administered was 100 mg of the test material.

6. The test material was ground to a fine powder prior to administration.

7. The eyes of the six animals in the nonwash group were not washed. The eyes of the three animals in the wash group were washed with lukewarm water for one minute no sooner than 30 seconds after treatment.

8. Both eyes of each animal were examined prior to treatment and only rabbits with normal ophthalmic findings were included in the test. The treated eyes of all animals were examined and evaluated for irritation at 1, 24, 48, and 72 hours and 4, 7, 10, and 13 days after treatment. Eyes of all nonwash animals were clear at Day 13, and eyes of all wash animals were clear at 72 hours. According to the eye irritation classification scheme, the nonwash treatment was moderately irritating and the wash treatment was minimally irritating.

9. Animals were not observed for pharmacological and/or toxicological effects on the day of dosing.

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10. Eye irritation evaluation in the nonwash animals beginning 24 hours after dosing found conjunctival redness, chemosis, and discharge clear by Day 10, and corneal opacity clear by Day 13. For the wash animals, all effects were clear by 72 hours. Average irritation scores for the nonwash animals were 20.3, 27.0, 25.0, 22.0, 7.5, 1.7, and 0.0 at 24, 28, and 72 hours and 4, 7, 10, and 13 days, respectively. Average irritation scores for the wash animals were 2.7, 0.7, and 0.0 at 24, 48, and 72 hours, respectively. Animals were not specifically observed for other pharmacological and/or toxicological effects for the 13 days following the day of dosing.

11. There were no significant deviations from the Acceptance Criteria in this study. Three deviations from the Acceptance Criteria are noted. Under Item 8, the eyes were not graded for irritation before dosing and each day after Day 4. This deviation is considered to be insignificant because the eyes were examined on Days 7, 10, and 13, and the irritation category would not be affected by changes occurring on intervening days. Under Items 9 and 10, individual observations for toxicity other than eye irritation were not recorded. These criteria are marked as supplemental. In addition, such observations would not substantially enhance the evaluation of potential eye irritation or potential systemic toxicity without the measurement of additional parameters.

GILLIS:R504SW0921JG/MT